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                  prophetic substances
 NEWS  4  JAN 28  USPATFULL, USPAT2, and USPATOLD enhanced with new
                  custom IPC display formats
 NEWS  5  JAN 28  MARPAT searching enhanced
 NEWS  6  JAN 28  USGENE now provides USPTO sequence data within 3 days
                  of publication
 NEWS  7  JAN 28  TOXCENTER enhanced with reloaded MEDLINE segment
 NEWS  8  JAN 28  MEDLINE and LMEDLINE reloaded with enhancements
 NEWS  9  FEB 08  STN Express, Version 8.3, now available
 NEWS 10  FEB 20  PCI now available as a replacement to DPCI
 NEWS 11  FEB 25  IFIREF reloaded with enhancements
 NEWS 12  FEB 25  IMSPRODUCT reloaded with enhancements
 NEWS 13  FEB 29  WPINDEX/WPIDS/WPIX enhanced with ECLA and current
                  U.S. National Patent Classification
 NEWS 14  MAR 31  IFICDB, IFIPAT, and IFIUDB enhanced with new custom
                  IPC display formats
 NEWS 15  MAR 31  CAS REGISTRY enhanced with additional experimental
                  spectra
 NEWS 16  MAR 31  CA/CAplus and CASREACT patent number format for U.S.
                  applications updated
 NEWS 17  MAR 31  LPCI now available as a replacement to LDPCI
 NEWS 18  MAR 31  EMBASE, EMBAL, and LEMBASE reloaded with enhancements
 NEWS 19  APR 04  STN AnaVist, Version 1, to be discontinued
 NEWS 20  APR 15  WPIDS, WPINDEX, and WPIX enhanced with new
                  predefined hit display formats
 NEWS 21  APR 28  EMBASE Controlled Term thesaurus enhanced
 NEWS 22  APR 28  IMSRESEARCH reloaded with enhancements

 NEWS EXPRESS FEBRUARY 08 CURRENT WINDOWS VERSION IS V8.3,
              AND CURRENT DISCOVER FILE IS DATED 20 FEBRUARY 2008

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* * * * * * * * * * * * * * * * STN  Columbus  * * * * * * * * * * * * * * * *

FILE 'HOME' ENTERED AT 19:46:42 ON 20 MAY 2008

=> file registry
COST IN U.S. DOLLARS                                    SINCE FILE      TOTAL
                                                           ENTRY    SESSION
FULL ESTIMATED COST                                         0.21       0.21

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DICTIONARY FILE UPDATES:  19 MAY 2008  HIGHEST RN 1021481-05-9

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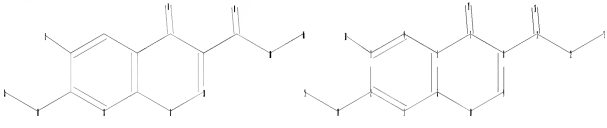

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ring nodes :
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2-14  3-18  7-17  8-11  11-12  11-16  12-13  14-15
ring bonds :
1-2  1-6  2-3  3-4  4-5  5-6  5-7  6-10  7-8  8-9  9-10
exact/norm bonds :
2-14  5-7  6-10  7-8  7-17  8-9  9-10  11-12  11-16  12-13
exact bonds :
3-18  8-11  14-15
normalized bonds :
1-2  1-6  2-3  3-4  4-5  5-6
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Match level :
1:Atom 2:Atom 3:Atom 4:Atom 5:Atom 6:Atom 7:Atom 8:Atom 9:Atom 10:Atom
11:CLASS 12:CLASS 13:CLASS 14:CLASS 15:CLASS 16:CLASS 17:CLASS 18:CLASS

L1 STRUCTURE UPLOADED

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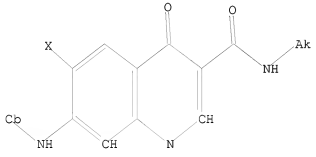
L1 HAS NO ANSWERS
L1 STR



Structure attributes must be viewed using STN Express query preparation.

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FULL SCREEN SEARCH COMPLETED - 3676 TO ITERATE

100.0% PROCESSED 3676 ITERATIONS 482 ANSWERS
SEARCH TIME: 00.00.01

L2 482 SEA SSS FUL L1

=> file caplus
COST IN U.S. DOLLARS SINCE FILE ENTRY TOTAL SESSION
FULL ESTIMATED COST 179.28 179.49

FILE 'CAPLUS' ENTERED AT 19:48:37 ON 20 MAY 2008
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FILE COVERS 1907 - 20 May 2008 VOL 148 ISS 21

FILE LAST UPDATED: 19 May 2008 (20080519/ED)

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=> s l2
L3 4 L2

=> d l3 1-4 abs ibib

L3 ANSWER 1 OF 4 CAPLUS COPYRIGHT 2008 ACS on STN
GI

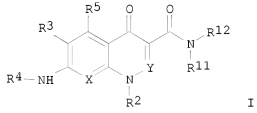
* STRUCTURE DIAGRAM TOO LARGE FOR DISPLAY - AVAILABLE VIA OFFLINE PRINT *

AB Process for producing compds. I [X = CR7, N; Y = CR6, N; R2 = (un)substituted alkyl, cycloalkyl, aryl, etc.; R3 = halo, alkyl, O-alkyl; R4 = (un)substituted cycloalkyl, non aromatic heterocycle, alkyl substituted by cycloalkyl; further detail on R4 is given.; R5 = H, halo, cyano, etc.; R6 = H, halo, alkyl, etc.; R7 = H, halo, alkyl, etc.; R11 = H, (un)substituted alkyl, optionally substituted amino by (un)substituted alkyl; R12 = H, (un)substituted alkyl, aryl; R11 and R12 may combine to form cyclic amino group in cooperation with the adjacent nitrogen.] or their pharmaceutically acceptable salts, characterized by reaction of compds. II [X, Y, R2-R5 = same as above] or active derivs. thereof with NHR11R12 [R11, R12 = same as above], was provided. For example, to a solution of compound III [R = OH; R' = cyclopentyl] (400 mg) in DMF (5.0 mL) was added 1,1'-carbonyldiimidazole (350 mg) at room temperature, the the reaction was stirred at 100 °C for 20 h. The resulting mixture was treated with Et3N (0.2 mL) and glycine Et ester hydrochloride (180 mg) at room temperature for 5 h to give compound III [R = NHCH2CO2Et; R' = cyclopentyl].
In platelet aggregation inhibition assays, compound III [R = NHCH2CH2P(:O)(OH)2; R' = 2,2-dimethyl-1,3-dioxan-5-yl] exhibited the activity of 92%.

ACCESSION NUMBER: 2006:882644 CAPLUS
DOCUMENT NUMBER: 145:292885
TITLE: Quinolone and related compounds as platelet aggregation inhibitors, and process for the preparation thereof
INVENTOR(S): Watanuki, Susumu; Koga, Yuji; Moritomo, Hiroyuki; Tsukamoto, Kazunari; Kaga, Daisuke; Okuda, Takao; Hirayama, Fukushi; Moritani, Yumiko; Takahashi, Atsushi
PATENT ASSIGNEE(S): Astellas Pharma Inc., Japan
SOURCE: Jpn. Kokai Tokkyo Koho, 95pp.
CODEN: JKXXAF
DOCUMENT TYPE: Patent
LANGUAGE: Japanese
FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
JP 2006225379	A	20060831	JP 2006-9367	20060118
PRIORITY APPLN. INFO.:			JP 2005-12618 A	20050120
OTHER SOURCE(S):	MARPAT 145:292885			

L3 ANSWER 2 OF 4 CAPLUS COPYRIGHT 2008 ACS on STN
GI



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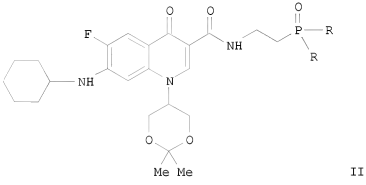

II

AB Title compds. I [X = CR7, N; Y = CR6, N; R11 = H, (un)substituted alkyl, optionally substituted amino by (un)substituted alkyl; R12 = H, (un)substituted alkyl, aryl; R11 and R12 may combine to form a (un)substituted cyclic amino group in cooperation with the adjacent nitrogen; R2 = (un)substituted alkyl, cycloalkyl, aryl, etc.; R3 = halo, alkyl, -O-alkyl; R4 = (un)substituted cycloalkyl, non aromatic heterocycle, alkyl substituted by cycloalkyl; further detail on R4 is given.; R5 = H, halo, cyano, etc.; R6 = H, halo, alkyl, etc.; R7 = H, halo, alkyl, etc.] and their pharmaceutically acceptable salts were prepared For example, Pd/C catalyzed debenzylation of compound II [R = OCH2Ph] under H2 afforded compound II [R = OH]. In platelet aggregation inhibition assays, compound II [R = OH] exhibited the activity of 92%.

ACCESSION NUMBER: 2006:882641 CAPLUS
DOCUMENT NUMBER: 145:292884
TITLE: Preparation of quinolone derivatives as platelet aggregation inhibitors
INVENTOR(S): Watanuki, Susumu; Koga, Yuji; Moritomo, Hiroyuki; Tsukamoto, Kazunari; Kaga, Daisuke; Okuda, Takao; Hirayama, Fukushi; Moritani, Yumiko; Takasaki, Atsushi
PATENT ASSIGNEE(S): Astellas Pharma Inc., Japan
SOURCE: Jpn. Kokai Tokkyo Koho, 95pp.
CODEN: JKXXAF
DOCUMENT TYPE: Patent
LANGUAGE: Japanese
FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE

JP 2006225378 A 20060831 JP 2006-9349 20060118
PRIORITY APPLN. INFO.: JP 2005-12561 A 20050120
OTHER SOURCE(S): MARPAT 145:292884

L3 ANSWER 3 OF 4 CAPLUS COPYRIGHT 2008 ACS on STN
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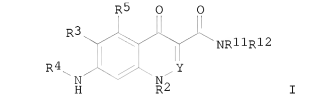

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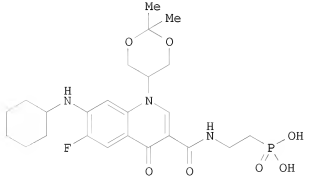


II

AB The title compds. (I) and pharmaceutically acceptable salts thereof characterized by each having an amide group at the 3-position which is substituted with a substituent having a carboxylate ester, phosphate ester, sulfate ester or the like, and an amino group at the 7-position which is substituted with a substituent having a ring structure [Y = C-R6; R6 = H, halo, lower alkyl, halo-lower alkyl; R2 = each (un)substituted lower alkyl, cycloalkyl, aryl, or heterocyclyl; R3 = halo; R5 = H, HO, halo; R11 = H, lower alkyl or lower alkyl-amino wherein lower alkyl is optionally substituted; R12 = (un)substituted lower alkyl] are prepared These compds. have excellent P2Y12 (adenine diphosphate receptor) inhibitory effect and platelet agglutination inhibitory effect and consequently are useful as platelet agglutination inhibitors. Thus, hydrogenolysis of [2-(([7-(Cyclohexylamino)-1-(2,2-dimethyl-1,3-dioxan-5-yl)-6-fluoro-4-oxo-1,4-dihydroquinolin-3-yl]carbonyl)amino)ethyl]phosphonic acid dibenzyl ester over 10% Pd-C in MeOH under hydrogen atmospheric for 3 h gave [2-(([7-(Cyclohexylamino)-1-(2,2-dimethyl-1,3-dioxan-5-yl)-6-fluoro-4-oxo-1,4-dihydroquinolin-3-yl]carbonyl)amino)ethyl]phosphonic acid (II). II inhibited ADP-induced aggregation of human blood platelet by 92% at 10 μM and the binding of [3H]-2-MeS-ADP to human P2Y12 by 96% at 30 nM.

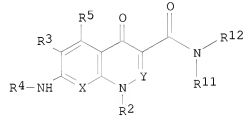
ACCESSION NUMBER: 2006:733081 CAPLUS
DOCUMENT NUMBER: 145:188746
TITLE: Preparation of 4-quinolone-3-carboxamide derivatives and salts thereof as platelet aggregation inhibitors
INVENTOR(S): Koga, Yuji; Okuda, Takao; Hirabayashi, Ryoji; Fujiyasu, Jiro; Miyazaki, Takehiro; Watanuki, Susumu; Hirayama, Fukushi; Moritani, Yumiko; Takasaki, Jun
PATENT ASSIGNEE(S): Astellas Pharma Inc., Japan
SOURCE: PCT Int. Appl., 150 pp.
CODEN: PIXXD2

DOCUMENT TYPE: Patent
LANGUAGE: Japanese
FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2006077851	A1	20060727	WO 2006-JP300590	20060118
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KM, KN, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, LY, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NG, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SM, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW				
RW: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LT, LU, LV, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG, BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
JP 2008094720	A	20080424	JP 2005-12715	20050120

PRIORITY APPLN. INFO.: JP 2005-12715 A 20050120
OTHER SOURCE(S): MARPAT 145:188746
REFERENCE COUNT: 6 THERE ARE 6 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L3 ANSWER 4 OF 4 CAPLUS COPYRIGHT 2008 ACS on STN
GI



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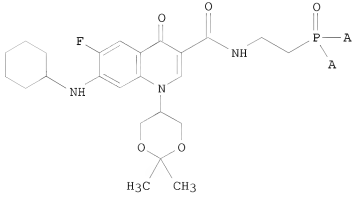

II

AB Title compds. I [X = CR7, N; Y = CR6, N; R11 = H, (un)substituted alkyl, etc.; R12 = H, (un)substituted alkyl, etc.; R2 = (un)substituted alkyl,

etc.; R3 = halo, etc.; R4 = (un)substituted cycloalkyl, etc.; R5 = H, halo, etc.; R6 = H, halo, etc.; R7 = H, halo, etc.] were prepared For example, hydrogenolysis of compound II [A = OCH2Ph] afforded compound II [A = OH]. In platelet aggregation inhibition assays, compound II [A = OH] exhibited inhibition activity of 92%. Compds. I are claimed useful as platelet aggregation inhibitors, P2Y12 inhibitors.

ACCESSION NUMBER: 2005:99478 CAPLUS
DOCUMENT NUMBER: 142:197896
TITLE: Preparation of quinolone derivatives as platelet aggregation inhibitors
INVENTOR(S): Watanuki, Susumu; Koga, Yuji; Moritomo, Hiroyuki; Tsukamoto, Issei; Kaga, Daisuke; Okuda, Takao; Hirayama, Fukushi; Moritani, Yumiko; Takasaki, Jun
PATENT ASSIGNEE(S): Yamanouchi Pharmaceutical Co., Ltd., Japan
SOURCE: PCT Int. Appl., 120 pp.
CODEN: PIXXD2
DOCUMENT TYPE: Patent
LANGUAGE: Japanese
FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2005009971	A1	20050203	WO 2004-JP10781	20040722
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RW: BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG				
JP 2005053903	A	20050303	JP 2004-212326	20040720
CA 2530352	A1	20050203	CA 2004-2530352	20040722
EP 1650192	A1	20060426	EP 2004-748045	20040722
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CN 1826321	A	20060830	CN 2004-80021187	20040722
US 20060148806	A1	20060706	US 2005-562128	20051223
IN 2006DN00144	A	20070824	IN 2006-DN144	20060109
MX 2006PA00675	A	20060419	MX 2006-PA675	20060118

PRIORITY APPLN. INFO.: JP 2003-278852 A 20030724
WO 2004-JP10781 W 20040722
OTHER SOURCE(S): MARPAT 142:197896
REFERENCE COUNT: 49 THERE ARE 49 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

=> file registry

COST IN U.S. DOLLARS	SINCE FILE ENTRY	TOTAL SESSION
FULL ESTIMATED COST	12.60	192.09

DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS)	SINCE FILE ENTRY	TOTAL SESSION
CA SUBSCRIBER PRICE	-3.20	-3.20

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STRUCTURE FILE UPDATES: 19 MAY 2008 HIGHEST RN 1021481-05-9
DICTIONARY FILE UPDATES: 19 MAY 2008 HIGHEST RN 1021481-05-9

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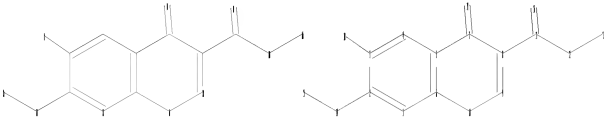

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ring nodes :
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2-14 3-18 7-17 8-11 11-12 11-16 12-13 14-15
ring bonds :
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exact/norm bonds :
2-14 5-7 6-10 7-8 7-17 8-9 9-10 11-12 11-16 12-13
exact bonds :
3-18 8-11 14-15
normalized bonds :
1-2 1-6 2-3 3-4 4-5 5-6

Match level :
1:Atom 2:Atom 3:Atom 4:Atom 5:Atom 6:Atom 7:Atom 8:Atom 9:Atom 10:Atom
11:CLASS 12:CLASS 13:CLASS 14:CLASS 15:CLASS 16:CLASS 17:CLASS 18:CLASS

L4   STRUCTURE UPLOADED

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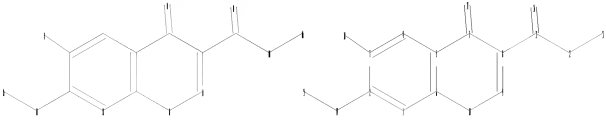

chain nodes :
11 12 13 14 15 16 17 18
ring nodes :
1 2 3 4 5 6 7 8 9 10
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ring bonds :
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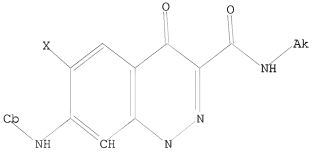
exact/norm bonds :
2-14 5-7 6-10 7-8 7-17 8-9 9-10 11-12 11-16 12-13
exact bonds :
3-18 8-11 14-15
normalized bonds :
1-2 1-6 2-3 3-4 4-5 5-6

Match level :
1:Atom 2:Atom 3:Atom 4:Atom 5:Atom 6:Atom 7:Atom 8:Atom 9:Atom 10:Atom
11:CLASS 12:CLASS 13:CLASS 14:CLASS 15:CLASS 16:CLASS 17:CLASS 18:CLASS

L5 STRUCTURE UPLOADED

=> d l5
L5 HAS NO ANSWERS
L5 STR



Structure attributes must be viewed using STN Express query preparation.

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FULL SCREEN SEARCH COMPLETED - 575 TO ITERATE

100.0% PROCESSED 575 ITERATIONS 13 ANSWERS
SEARCH TIME: 00.00.01

L6 13 SEA SSS FUL L5

=> file caplus

COST IN U.S. DOLLARS	SINCE FILE ENTRY	TOTAL SESSION
FULL ESTIMATED COST	178.82	370.91

DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS)	SINCE FILE ENTRY	TOTAL SESSION
CA SUBSCRIBER PRICE	0.00	-3.20

FILE 'CAPLUS' ENTERED AT 19:51:05 ON 20 MAY 2008
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L7 4 L6

=> d l7 1-4 abs ibib

L7 ANSWER 1 OF 4 CAPLUS COPYRIGHT 2008 ACS on STN
GI

* STRUCTURE DIAGRAM TOO LARGE FOR DISPLAY - AVAILABLE VIA OFFLINE PRINT *

AB Process for producing compds. I [X = CR7, N; Y = CR6, N; R2 = (un)substituted alkyl, cycloalkyl, aryl, etc.; R3 = halo, alkyl, O-alkyl; R4 = (un)substituted cycloalkyl, non aromatic heterocycle, alkyl substituted by cycloalkyl; further detail on R4 is given.; R5 = H, halo, cyano, etc.; R6 = H, halo, alkyl, etc.; R7 = H, halo, alkyl, etc.; R11 = H, (un)substituted alkyl, optionally substituted amino by (un)substituted alkyl; R12 = H, (un)substituted alkyl, aryl; R11 and R12 may combine to form cyclic amino group in cooperation with the adjacent nitrogen.] or their pharmaceutically acceptable salts, characterized by reaction of compds. II [X, Y, R2-R5 = same as above] or active derivs. thereof with NHR11R12 [R11, R12 = same as above], was provided. For example, to a

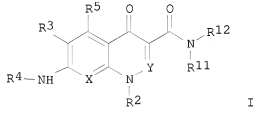
solution of compound III [R = OH; R' = cyclopentyl] (400 mg) in DMF (5.0 mL) was added 1,1'-carbonyldiimidazole (350 mg) at room temperature, the the reaction was stirred at 100 °C for 20 h. The resulting mixture was treated with Et3N (0.2 mL) and glycine Et ester hydrochloride (180 mg) at room temperature for 5 h to give compound III [R = NHCH2CO2Et; R' = cyclopentyl].

In platelet aggregation inhibition assays, compound III [R = NHCH2CH2P(:O)(OH)2; R' = 2,2-dimethyl-1,3-dioxan-5-yl] exhibited the activity of 92%.

ACCESSION NUMBER: 2006:882644 CAPLUS
DOCUMENT NUMBER: 145:292885
TITLE: Quinolone and related compounds as platelet aggregation inhibitors, and process for the preparation thereof
INVENTOR(S): Watanuki, Susumu; Koga, Yuji; Moritomo, Hiroyuki; Tsukamoto, Kazunari; Kaga, Daisuke; Okuda, Takao; Hirayama, Fukushi; Moritani, Yumiko; Takahashi, Atsushi
PATENT ASSIGNEE(S): Astellas Pharma Inc., Japan
SOURCE: Jpn. Kokai Tokkyo Koho, 95pp.
CODEN: JKXXAF
DOCUMENT TYPE: Patent
LANGUAGE: Japanese
FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
JP 2006225379	A	20060831	JP 2006-9367	20060118
PRIORITY APPLN. INFO.:			JP 2005-12618 A	20050120
OTHER SOURCE(S):	MARPAT 145:292885			

L7 ANSWER 2 OF 4 CAPLUS COPYRIGHT 2008 ACS on STN
GI



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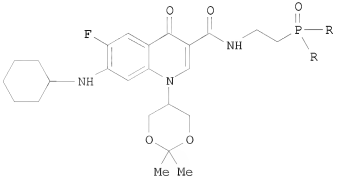

II

AB Title compds. I [X = CR7, N; Y = CR6, N; R11 = H, (un)substituted alkyl, optionally substituted amino by (un)substituted alkyl; R12 = H, (un)substituted alkyl, aryl; R11 and R12 may combine to form a (un)substituted cyclic amino group in cooperation with the adjacent nitrogen; R2 = (un)substituted alkyl, cycloalkyl, aryl, etc.; R3 = halo, alkyl, -O-alkyl; R4 = (un)substituted cycloalkyl, non aromatic heterocycle, alkyl substituted by cycloalkyl; further detail on R4 is given.; R5 = H, halo, cyano, etc.; R6 = H, halo, alkyl, etc.; R7 = H, halo, alkyl, etc.] and their pharmaceutically acceptable salts were prepared For example, Pd/C catalyzed debenzylation of compound II [R = OCH2Ph] under H2 afforded compound II [R = OH]. In platelet aggregation inhibition assays, compound II [R = OH] exhibited the activity of 92%.

ACCESSION NUMBER: 2006:882641 CAPLUS
DOCUMENT NUMBER: 145:292884
TITLE: Preparation of quinolone derivatives as platelet aggregation inhibitors
INVENTOR(S): Watanuki, Susumu; Koga, Yuji; Moritomo, Hiroyuki; Tsukamoto, Kazunari; Kaga, Daisuke; Okuda, Takao; Hirayama, Fukushi; Moritani, Yumiko; Takasaki, Atsushi
PATENT ASSIGNEE(S): Astellas Pharma Inc., Japan
SOURCE: Jpn. Kokai Tokkyo Koho, 95pp.
CODEN: JKXXAF
DOCUMENT TYPE: Patent
LANGUAGE: Japanese
FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.		DATE
JP 2006225378	A	20060831	JP 2006-9349		20060118
PRIORITY APPLN. INFO.:			JP 2005-12561	A	20050120
OTHER SOURCE(S):	MARPAT 145:292884				

L7 ANSWER 3 OF 4 CAPLUS COPYRIGHT 2008 ACS on STN
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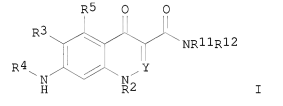

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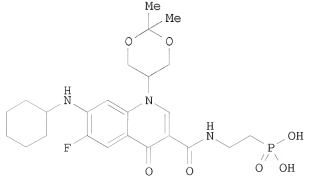

II

AB The title compds. (I) and pharmaceutically acceptable salts thereof characterized by each having an amide group at the 3-position which is substituted with a substituent having a carboxylate ester, phosphate ester, sulfate ester or the like, and an amino group at the 7-position which is substituted with a substituent having a ring structure [Y = C-R6; R6 = H, halo, lower alkyl, halo-lower alkyl; R2 = each (un)substituted lower alkyl, cycloalkyl, aryl, or heterocyclyl; R3 = halo; R5 = H, HO, halo; R11 = H, lower alkyl or lower alkyl-amino wherein lower alkyl is optionally substituted; R12 = (un)substituted lower alkyl] are prepared These compds. have excellent P2Y12 (adenine diphosphate receptor) inhibitory effect and platelet agglutination inhibitory effect and consequently are useful as platelet agglutination inhibitors. Thus, hydrogenolysis of [2-(([7-(Cyclohexylamino)-1-(2,2-dimethyl-1,3-dioxan-5-yl)-6-fluoro-4-oxo-1,4-dihydroquinolin-3-yl]carbonyl)amino)ethyl]phosphonic acid dibenzyl ester over 10% Pd-C in MeOH under hydrogen atmospheric for 3 h gave [2-(([7-(Cyclohexylamino)-1-(2,2-dimethyl-1,3-dioxan-5-yl)-6-fluoro-4-oxo-1,4-dihydroquinolin-3-yl]carbonyl)amino)ethyl]phosphonic acid (II). II inhibited ADP-induced aggregation of human blood platelet by 92% at 10 μM and the binding of [3H]-2-MeS-ADP to human P2Y12 by 96% at 30 nM.

ACCESSION NUMBER: 2006:733081 CAPLUS
DOCUMENT NUMBER: 145:188746
TITLE: Preparation of 4-quinolone-3-carboxamide derivatives and salts thereof as platelet aggregation inhibitors
INVENTOR(S): Koga, Yuji; Okuda, Takao; Hirabayashi, Ryoji; Fujiyasu, Jiro; Miyazaki, Takehiro; Watanuki, Susumu; Hirayama, Fukushi; Moritani, Yumiko; Takasaki, Jun
PATENT ASSIGNEE(S): Astellas Pharma Inc., Japan
SOURCE: PCT Int. Appl., 150 pp.
CODEN: PIXXD2
DOCUMENT TYPE: Patent
LANGUAGE: Japanese
FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2006077851	A1	20060727	WO 2006-JP300590	20060118
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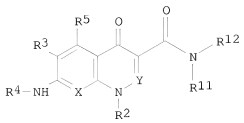
JP 2008094720	A	20080424	JP 2005-12715	20050120
PRIORITY APPLN. INFO.:			JP 2005-12715 A	20050120
OTHER SOURCE(S):	MARPAT 145:188746			
REFERENCE COUNT:	6	THERE ARE 6 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT		

L7 ANSWER 4 OF 4 CAPLUS COPYRIGHT 2008 ACS on STN
GI



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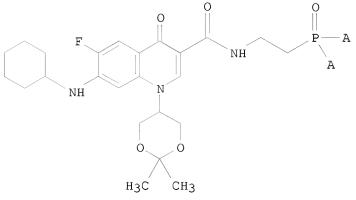

II

AB Title compds. I [X = CR7, N; Y = CR6, N; R11 = H, (un)substituted alkyl, etc.; R12 = H, (un)substituted alkyl, etc.; R2 = (un)substituted alkyl, etc.; R3 = halo, etc.; R4 = (un)substituted cycloalkyl, etc.; R5 = H, halo, etc.; R6 = H, halo, etc.; R7 = H, halo, etc.] were prepared For example, hydrogenolysis of compound II [A = OCH2Ph] afforded compound II [A = OH]. In platelet aggregation inhibition assays, compound II [A = OH] exhibited inhibition activity of 92%. Compds. I are claimed useful as platelet aggregation inhibitors, P2Y12 inhibitors.

ACCESSION NUMBER: 2005:99478 CAPLUS
DOCUMENT NUMBER: 142:197896
TITLE: Preparation of quinolone derivatives as platelet aggregation inhibitors
INVENTOR(S): Watanuki, Susumu; Koga, Yuji; Moritomo, Hiroyuki; Tsukamoto, Issei; Kaga, Daisuke; Okuda, Takao; Hirayama, Fukushi; Moritani, Yumiko; Takasaki, Jun
PATENT ASSIGNEE(S): Yamanouchi Pharmaceutical Co., Ltd., Japan
SOURCE: PCT Int. Appl., 120 pp.
CODEN: PIXXD2
DOCUMENT TYPE: Patent
LANGUAGE: Japanese
FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2005009971	A1	20050203	WO 2004-JP10781	20040722
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     JP 2005053903          A     20050303    JP 2004-212326          20040720
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     EP 1650192             A1    20060426    EP 2004-748045          20040722
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     CN 1826321             A     20060830    CN 2004-80021187        20040722
     US 20060148806         A1    20060706    US 2005-562128          20051223
     IN 2006DN00144         A     20070824    IN 2006-DN144           20060109
     MX 2006PA00675         A     20060419    MX 2006-PA675           20060118
PRIORITY APPLN. INFO.:                         JP 2003-278852      A  20030724
                                               WO 2004-JP10781     W  20040722
OTHER SOURCE(S):          MARPAT 142:197896
REFERENCE COUNT:          49     THERE ARE 49 CITED REFERENCES AVAILABLE FOR THIS
                                 RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT


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COST IN U.S. DOLLARS                                SINCE FILE      TOTAL
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FULL ESTIMATED COST                                    12.12     383.03

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DICTIONARY FILE UPDATES:  19 MAY 2008  HIGHEST RN 1021481-05-9

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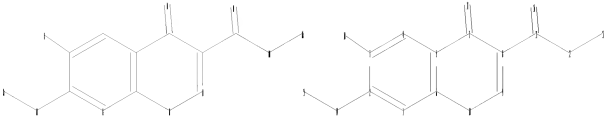

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ring bonds :
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normalized bonds :
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L8      STRUCTURE UPLOADED

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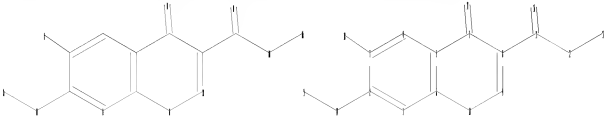

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chain nodes :
11  12  13  14  15  16  17  18
ring nodes :
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1  2  3  4  5  6  7  8  9  10
chain bonds :
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ring bonds :
1-2  1-6  2-3  3-4  4-5  5-6  5-7  6-10  7-8  8-9  9-10
exact/norm bonds :
2-14  5-7  6-10  7-8  7-17  8-9  9-10  11-12  11-16  12-13
exact bonds :
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normalized bonds :
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Match level :
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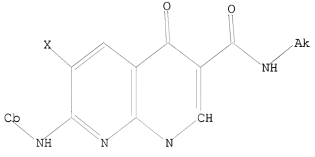


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FILE COVERS 1907 - 20 May 2008 VOL 148 ISS 21
FILE LAST UPDATED: 19 May 2008 (20080519/ED)

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L12 3 L11

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L12 ANSWER 1 OF 3 CAPLUS COPYRIGHT 2008 ACS on STN
GI

* STRUCTURE DIAGRAM TOO LARGE FOR DISPLAY - AVAILABLE VIA OFFLINE PRINT *

AB Process for producing compds. I [X = CR7, N; Y = CR6, N; R2 = (un)substituted alkyl, cycloalkyl, aryl, etc.; R3 = halo, alkyl, O-alkyl; R4 = (un)substituted cycloalkyl, non aromatic heterocycle, alkyl substituted by cycloalkyl; further detail on R4 is given.; R5 = H, halo, cyano, etc.; R6 = H, halo, alkyl, etc.; R7 = H, halo, alkyl, etc.; R11 = H, (un)substituted alkyl, optionally substituted amino by (un)substituted alkyl; R12 = H, (un)substituted alkyl, aryl; R11 and R12 may combine to form cyclic amino group in cooperation with the adjacent nitrogen.] or their pharmaceutically acceptable salts, characterized by reaction of compds. II [X, Y, R2-R5 = same as above] or active derivs. thereof with NHR11R12 [R11, R12 = same as above], was provided. For example, to a solution of compound III [R = OH; R' = cyclopentyl] (400 mg) in DMF (5.0 mL) was added 1,1'-carbonyldiimidazole (350 mg) at room temperature, the the reaction was stirred at 100 °C for 20 h. The resulting mixture was treated with Et3N (0.2 mL) and glycine Et ester hydrochloride (180 mg) at room temperature for 5 h to give compound III [R = NHCH2CO2Et; R' = cyclopentyl].
In platelet aggregation inhibition assays, compound III [R = NHCH2CH2P(:O)(OH)2; R' = 2,2-dimethyl-1,3-dioxan-5-yl] exhibited the activity of 92%.

ACCESSION NUMBER: 2006:882644 CAPLUS
DOCUMENT NUMBER: 145:292885
TITLE: Quinolone and related compounds as platelet aggregation inhibitors, and process for the preparation thereof
INVENTOR(S): Watanuki, Susumu; Koga, Yuji; Moritomo, Hiroyuki;

Tsukamoto, Kazunari; Kaga, Daisuke; Okuda, Takao; Hirayama, Fukushi; Moritani, Yumiko; Takahashi, Atsushi
PATENT ASSIGNEE(S): Astellas Pharma Inc., Japan
SOURCE: Jpn. Kokai Tokkyo Koho, 95pp.
CODEN: JKXXAF
DOCUMENT TYPE: Patent
LANGUAGE: Japanese
FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
JP 2006225379	A	20060831	JP 2006-9367	20060118
PRIORITY APPLN. INFO.:			JP 2005-12618 A	20050120

OTHER SOURCE(S): MARPAT 145:292885

L12 ANSWER 2 OF 3 CAPLUS COPYRIGHT 2008 ACS on STN
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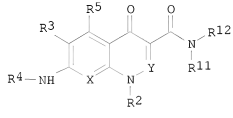

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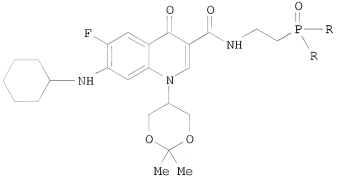

II

AB Title compds. I [X = CR7, N; Y = CR6, N; R11 = H, (un)substituted alkyl, optionally substituted amino by (un)substituted alkyl; R12 = H, (un)substituted alkyl, aryl; R11 and R12 may combine to form a (un)substituted cyclic amino group in cooperation with the adjacent nitrogen; R2 = (un)substituted alkyl, cycloalkyl, aryl, etc.; R3 = halo, alkyl, -O-alkyl; R4 = (un)substituted cycloalkyl, non aromatic heterocycle, alkyl substituted by cycloalkyl; further detail on R4 is given.; R5 = H, halo, cyano, etc.; R6 = H, halo, alkyl, etc.; R7 = H, halo, alkyl, etc.] and their pharmaceutically acceptable salts were prepared For example, Pd/C catalyzed debenzylation of compound II [R = OCH2Ph] under H2 afforded compound II [R = OH]. In platelet aggregation inhibition assays, compound II [R = OH] exhibited the activity of 92%.
ACCESSION NUMBER: 2006:882641 CAPLUS

DOCUMENT NUMBER: 145:292884
TITLE: Preparation of quinolone derivatives as platelet aggregation inhibitors
INVENTOR(S): Watanuki, Susumu; Koga, Yuji; Moritomo, Hiroyuki; Tsukamoto, Kazunari; Kaga, Daisuke; Okuda, Takao; Hirayama, Fukushi; Moritani, Yumiko; Takasaki, Atsushi
PATENT ASSIGNEE(S): Astellas Pharma Inc., Japan
SOURCE: Jpn. Kokai Tokkyo Koho, 95pp.
CODEN: JKXXAF
DOCUMENT TYPE: Patent
LANGUAGE: Japanese
FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
JP 2006225378	A	20060831	JP 2006-9349	20060118
PRIORITY APPLN. INFO.:			JP 2005-12561 A	20050120
OTHER SOURCE(S):		MARPAT 145:292884		

L12 ANSWER 3 OF 3 CAPLUS COPYRIGHT 2008 ACS on STN
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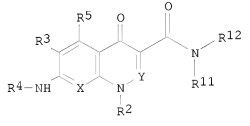
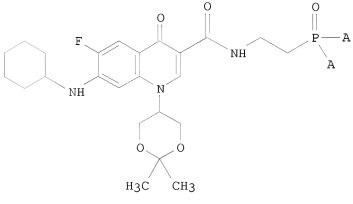


I



II

AB Title compds. I [X = CR7, N; Y = CR6, N; R11 = H, (un)substituted alkyl, etc.; R12 = H, (un)substituted alkyl, etc.; R2 = (un)substituted alkyl, etc.; R3 = halo, etc.; R4 = (un)substituted cycloalkyl, etc.; R5 = H, halo, etc.; R6 = H, halo, etc.; R7 = H, halo, etc.] were prepared For example, hydrogenolysis of compound II [A = OCH2Ph] afforded compound II [A = OH]. In platelet aggregation inhibition assays, compound II [A = OH] exhibited inhibition activity of 92%. Compds. I are claimed useful as platelet aggregation inhibitors, P2Y12 inhibitors.

ACCESSION NUMBER: 2005:99478 CAPLUS
DOCUMENT NUMBER: 142:197896
TITLE: Preparation of quinolone derivatives as platelet aggregation inhibitors
INVENTOR(S): Watanuki, Susumu; Koga, Yuji; Moritomo, Hiroyuki; Tsukamoto, Issei; Kaga, Daisuke; Okuda, Takao; Hirayama, Fukushi; Moritani, Yumiko; Takasaki, Jun
PATENT ASSIGNEE(S): Yamanouchi Pharmaceutical Co., Ltd., Japan
SOURCE: PCT Int. Appl., 120 pp.
CODEN: PIXXD2
DOCUMENT TYPE: Patent
LANGUAGE: Japanese
FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2005009971	A1	20050203	WO 2004-JP10781	20040722
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW				
RW: BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG				
JP 2005053903	A	20050303	JP 2004-212326	20040720
CA 2530352	A1	20050203	CA 2004-2530352	20040722
EP 1650192	A1	20060426	EP 2004-748045	20040722
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, FI, RO, CY, TR, BG, CZ, EE, HU, PL, SK				
CN 1826321	A	20060830	CN 2004-80021187	20040722
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IN 2006DN00144	A	20070824	IN 2006-DN144	20060109
MX 2006PA00675	A	20060419	MX 2006-PA675	20060118
PRIORITY APPLN. INFO.:			JP 2003-278852	A 20030724
			WO 2004-JP10781	W 20040722
OTHER SOURCE(S):	MARPAT 142:197896			
REFERENCE COUNT:	49	THERE ARE 49 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT		

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